# PATENT COOPERATION TREATY

# **PCT**

# INTERNATIONAL SEARCH REPORT

(PCT Article 18 and Rules 43 and 44)

Applicant's or agent's file reference FOR FURTHER see Notification of Transmittal of International Search Report (Form PCT/ISA/220) as well as, where applicable, item 5 below.						
SH/TC/34953 ACTION						
International application No.	national application No. International filing date (day/month/year) (Earliest) Priority Date (day/month/year)					
PCT/IB 99/02018	26/11/1999	27/11/1998				
Applicant						
LUDUTO INSTITUTE FOR CANOL	ED DECEADON -+ -1					
LUDWIG INSTITUTE FOR CANCI	ER RESEARCH et al.					
This lutery stirms I Seems Bonort has been	n prepared by this International Searching Auth	pority and is transmitted to the applicant				
according to Article 18. A copy is being tra		ionly and is transmitted to the applicant				
This International Search Report consists  X It is also accompanied by	of a total of sheets.  a copy of each prior art document cited in this	report.				
Basis of the report      With regard to the language the	international search was carried out on the ba	sis of the international application in the				
	ess otherwise indicated under this item.	sis of the international approach in the				
the international search w Authority (Rule 23.1(b)).	as carried out on the basis of a translation of t	he international application furnished to this				
		nternational application, the international search				
was carried out on the basis of the X contained in the internation	e sequence listing . nal application in written form.	•				
T filed together with the inte	rnational application in computer readable for	n.				
furnished subsequently to	this Authority in written form.					
1 🗎	this Authority in computer readble form.					
	osequently furnished written sequence listing d s filed has been furnished.	loes not go beyond the disclosure in the				
the statement that the info furnished	ormation recorded in computer readable form i	s identical to the written sequence listing has been				
2. X Certain claims were fou	nd unsearchable (See Box I).					
3. X Unity of invention is lac	king (see Box II).					
4. With regard to the title,						
the text is approved as su	bmitted by the applicant.					
	hed by this Authority to read as follows:					
MAGE-10 OR MAGE-8 DERIVED HLA-A2.1-BINDING OLIGOPEPTIDES						
5. With regard to the <b>abstract,</b>						
X the text is approved as submitted by the applicant.						
the text has been established, according to Rule 38.2(b), by this Authority as it appears in Box III. The applicant may, within one month from the date of mailing of this international search report, submit comments to this Authority.						
6. The figure of the <b>drawings</b> to be published.						
as suggested by the appl		None of the figures.				
because the applicant fai	ed to suggest a figure.					
because this figure better	characterizes the invention.					

International application No. PCT/IB 99/02018 1/

# INTERNATIONAL SEARCH REPORT

Box I Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)
This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1. X Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:
Although claim is 39 directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.
2. X Claims Nos.: - because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
see FURTHER INFORMATION sheet PCT/ISA/210
3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)
This International Searching Authority found multiple inventions in this international application, as follows:
see additional sheet
As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4. No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:
Remark on Protest  The additional search fees were accompanied by the applicant's protest.  X  No protest accompanied the payment of additional search fees.

# FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

This International Searching Authority found multiple (groups of) inventions in this international application, as follows:

1. Claims: 1-11 and 13-39 partially, and 12 completely

Polypeptides comprising an unbroken sequence from seq.ID.1, either capabale of binding HLA-A2 or of eliciting an immune response from human lymphocytes, nucleic acids encoding them, expression vectors comprising said nucleic acids, hosts comprising said vectors, agents capable of specifically binding said polypeptides, pharmaceutical compositions of said polypeptides, optionally in combination with said HLA, said vector, said host or said binding agent, a cell pulsed with said polypetide presenting it on said HLA on its surface, method of diagnosis using said binding aganet or said nucleic acid, and a method of producing a CTL culture using said polypeptide and said HLA.

Seq.ID's 43 and 45, derived from seq.ID.2, have been searched with this invention due to their similarity with seq.ID's 42 and 44, respectively.

2. Claims: 1-11, 13-39, all partially

Polypeptides comprising an unbroken sequence from seq.ID.2, either capabale of binding HLA-A2 or of eliciting an immune response from human lymphocytes, nucleic acids encoding them, expression vectors comprising said nucleic acids, hosts comprising said vectors, agents capable of specifically binding said polypeptides, pharmaceutical compositions of said polypeptides, optionally in combination with said HLA, said vector, said host or said binding agent, a cell pulsed with said polypetide presenting it on said HLA on its surface, method of diagnosis using said binding aganet or said nucleic acid, and a method of producing a CTL culture using said polypeptide and said HLA.

Seq.ID's 43 and 45 have been searched with the first invention.

# FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

#### Continuation of Box I.2

Present claims 1,2,5-7, and 10-39 relate to an extremely large number of possible polypeptides, defined by reference to a desirable characteristic or property, namely their ability to form a complex with an HLA-A2 type molecule and/ or their ability to elicit an immune response. The claims contain so many possible permutations that a lack of clarity (and/or conciseness) within the meaning of Article 6 PCT arises. Furthermore, the claims cover all polypeptides having the desired characteristic or property, whereas the application provides support within the meaning of Article 6 PCT and/or disclosure within the meaning of Article 5 PCT for only a very limited number of such polypeptides. In the present case, the claims so lack support, and the application so lacks disclosure, clarity and conciseness, that a meaningful search over the whole of the claimed scope is impossible. Furthermore, the claims contain so many possible permutations that a lack of clarity (and/or conciseness) within the meaning of Article 6 PCT arises. Independent of the above reasoning, the claims also lack clarity (Article 6 PCT). An attempt is made to define the polypeptides by reference to a result to be achieved. Again, this lack of clarity in the present case is such as to render a meaningful search over the whole of the claimed scope impossible. Consequently, the search has been carried out for those parts of the claims which appear to be clear, supported and disclosed, namely those parts relating to the nonapeptides defined by claim 3, and larger polypeptides comprising said nonapetidic sequences, e.g. the decapeptides of claim 9.

The applicant's attention is drawn to the fact that claims, or parts of claims, relating to inventions in respect of which no international search report has been established need not be the subject of an international preliminary examination (Rule 66.1(e) PCT). The applicant is advised that the EPO policy when acting as an International Preliminary Examining Authority is normally not to carry out a preliminary examination on matter which has not been searched. This is the case irrespective of whether or not the claims are amended following receipt of the search report or during any Chapter II procedure.

International Application No PCT/IB 99/02018

A. CLASSIFICATION OF SUBJECT MATTER
IPC 7 C12N15/12 C07K14/47

A61K31/70

A61K38/08

C07K14/705 A61K35/14

C07K7/06 A61K38/17 C07K16/18 G01N33/53

According to International Patent Classification (IPC) or to both national classification and IPC

### B. FIELDS SEARCHED

 $\begin{array}{ll} \mbox{Minimum documentation searched} & \mbox{(classification system followed by classification symbols)} \\ \mbox{IPC 7} & \mbox{C07K} & \mbox{C12N} & \mbox{A61K} & \mbox{G01N} \end{array}$ 

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

C. DOCUMENTS CONSIDERED TO BE RELEVANT						
Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.				
X	US 5 686 068 A (BOON-FALLEUR THIERRY ET AL) 11 November 1997 (1997-11-11)	1-3,5-7, 10,11, 13-20, 23,29,30				
	seq.ID's 31,32 and 57 and the claims tables 1,2					
X	VISSEREN, M.J.W. ET AL.: "Identification of HLA-A*0201-resticted CTL epitopes encoded by the tumor-specific MAGE-2 gene product."  INTERNATIONAL JOURNAL OF CANCER, vol. 73, no. 1, 26 September 1997 (1997-09-26), pages 125-30, XP000914539 table 1, peptide entry under "M2 181-189"	1-3,5,6, 10,11, 13-21, 23,29,30				
	- <b>/-</b> -					

X Further documents are listed in the continuation of box C.	Patent family members are listed in annex.
"A" document defining the general state of the art which is not considered to be of particular relevance "E" earlier document but published on or after the International filing date  "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)  "O" document referring to an oral disclosure, use, exhibition or other means  "P" document published prior to the international filing date but later than the priority date claimed	<ul> <li>"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention</li> <li>"X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone</li> <li>"Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.</li> <li>"&amp;" document member of the same patent family</li> </ul>
Date of the actual completion of the international search  8 June 2000	Date of mailing of the international search report  2.7. 06. 00
Name and mailing address of the ISA  European Patent Office, P.B. 5818 Patentlaan 2  NL – 2280 HV Rijswijk  Tel. (+31-70) 340-2040, Tx. 31 651 epo nl,  Fax: (+31-70) 340-3016	Authorized officer  Smalt, R

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International Application No
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C.(Continua	ation) DOCUMENTS CONSIDERED TO BE RELEVANT	
Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Х	CHEN, YT. ET AL.: "Identification of multiple cancer/testis antigens by allogenic antibody screening of a melanoma cell line library." PROC.NAT'L.ACAD.SCI.USA, vol. 95, June 1998 (1998-06), pages 6919-23, XP002132946 page 6921, right-hand column, paragraph 3 - paragraph 4; figure 2	1,2,5,6, 12-18, 20,21,23
X	WO 92 20356 A (LUDWIG INST CANCER RES) 26 November 1992 (1992-11-26)  whole document, particularly seq.ID.'s 9,20 and 22 and the claims. claims 47,56,101,123,126	1,2,5, 10-26, 29,30, 34,35,39
X	WO 98 14463 A (LUDWIG INST CANCER RES) 9 April 1998 (1998-04-09)	2,12,13, 15-17, 25,26, 29,34, 35,39
	see the whole document, particularly example 11. the whole document	
A	RAMMENSEE H -G ET AL: "MHC LIGANDS AND PEPTIDE MOTIFS: FIRST LISTING" IMMUNOGENETICS, DE, SPRINGER VERLAG, BERLIN, vol. 41, no. 4, 1 February 1995 (1995-02-01), pages 178-228, XP000673045 ISSN: 0093-7711 cited in the application the whole document	3
A	WO 95 25530 A (UNIV LEIDEN ;LUDWIG INST CANCER RES (US)) 28 September 1995 (1995-09-28) the whole document	
Ρ,Χ	HUANG, LQ. ET AL.: "Cytolytic T lymphocytes recognize an antigen encoded by MAGE-A10 on human melanoma." JOURNAL OF IMMUNOLOGY, vol. 162, 1 June 1999 (1999-06-01), pages 6849-54, XP002132947 the whole document	1-39
P,X	WO 99 54738 A (LUDWIG INST CANCER RES) 28 October 1999 (1999-10-28) see seq.ID.1 and claims.	1,2,5,6, 12-18, 20,21,23
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International Application No
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Category °	ation) DOCUMENTS CONSIDERED TO BE RELEVANT  Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Jategory *	Citation of document, with indication, where appropriate, of the relevant passages	Treievant to Classiff NO.
Р,Х	WO 99 45954 A (EPIMMUNE INC) 16 September 1999 (1999-09-16) page 95, 4th row of the table; page 102,	1-3,6,7, 10,11, 13-17
Ρ,Χ	12th row of the table.  RIMOLDI, D. ET AL.: "cDNA and protein characterization of human MAGE-10." INTERNATIONAL JOURNAL OF CANCER, vol. 82, no. 6, 9 September 1999 (1999-09-09), pages 901-7, XP000877098 page 902, right-hand column, paragraph 2 page 907, left-hand column, paragraph 2	2,12,13, 15-17, 25,26, 29,34, 35,39
E	WO 99 61916 A (SIDNEY JOHN ;EPIMMUNE INC (US); SOUTHWOOD SCOTT (US); SETTE ALESSA) 2 December 1999 (1999-12-02) see page 42, peptide 39.0390, and page 43, peptide 39.0403	1,2,5,6, 10-17, 25,30

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Information on patent family members

International Application No
PCT/IB 99/02018

Patent documen cited in search rep		Publication date		atent family nember(s)	Publication date
US 5686068	A	11-11-1997	US AU CA CN EP WO ZA AU CA CN EP FI JP NO NZ WO US ZA	5554724 A 710887 B 4045397 A 2261579 A 1226897 A 0964868 A 9804582 A 9706476 A 682597 B 2189395 A 2186006 A 1149257 A 0810870 A 963780 A 10502329 T 963918 A 283561 A 9525530 A 6063900 A 9502375 A	10-09-1996 30-09-1999 20-02-1998 05-02-1998 25-08-1999 22-12-1999 05-02-1998 01-09-1998 09-10-1997 09-10-1995 28-09-1995 07-05-1997 10-12-1997 23-09-1996 03-03-1998 20-11-1996 27-04-1998 28-09-1995 16-05-2000 15-12-1995
WO 9220356	Α	26-11-1992	US AU CA EP FI JP NO NZ PT US US US US US	5342774 A 664560 B 2158392 A 2109727 A 0595838 A 935174 A 6511144 T 934130 A 242875 A 100515 A,B 5925729 A 5541104 A 5612201 A 5843448 A 6025474 A 9203759 A	30-08-1994 23-11-1995 30-12-1992 26-11-1992 11-05-1994 22-11-1993 15-12-1994 23-11-1993 27-09-1993 31-08-1993 20-07-1999 30-07-1996 18-03-1997 01-12-1998 15-02-2000 28-04-1993
WO 9814463	A	09-04-1998	US AU CN EP ZA	5908778 A 4264097 A 1239479 A 0948517 A 9708802 A	01-06-1999 24-04-1998 22-12-1999 13-10-1999 09-02-1999
WO 9525530	<b>A</b>	28-09-1995	US AU CA CN EP FI JP NO NZ US US ZA	5554724 A 682597 B 2189395 A 2186006 A 1149257 A 0810870 A 963780 A 10502329 T 963918 A 283561 A 6063900 A 5686068 A 9502375 A	10-09-1996 09-10-1997 09-10-1995 28-09-1995 07-05-1997 10-12-1997 23-09-1996 03-03-1998 20-11-1996 27-04-1998 16-05-2000 11-11-1997 15-12-1995

Information on patent family members

International Application No
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Patent document cited in search report		Publication date	Patent family member(s)		Publication date
WO 9954738	Α	28-10-1999	AU	3007999 A	08-11-1999
WO 9945954	Α	16-09-1999	AU	6465598 A	27-09-1999
WO 9961916	Α	02-12-1999	AU	4224499 A	13-12-1999

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